

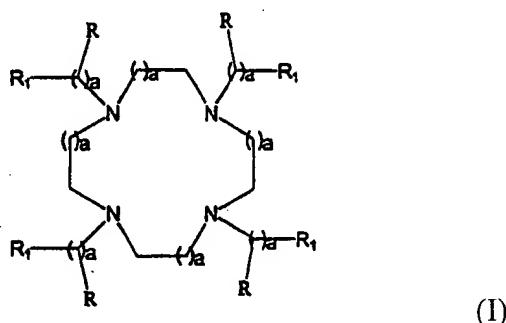
Claims (clean version encompassing amendments)



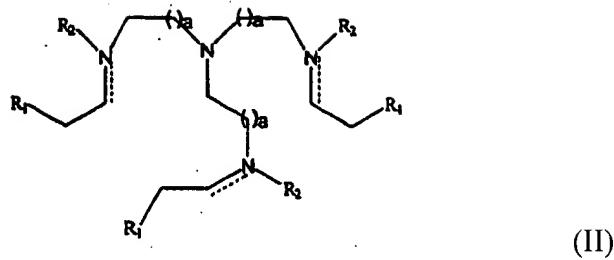
(twice amended) The method as claimed in claim 24, wherein said chelate complex is a complex of a chelant selected from the group consisting of DTPA, EDTA, DTPA-BMA, DO3A, DOTA, HP-DO3A, TMT and DPDP.

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11. (twice amended) The method as claimed in claim 24, wherein said chelate complex is a complex of a chelant selected from the group consisting of porphyrins, phthalocyanins, crown ethers, hemin, heme, chelants having a square planar symmetry, cryptands and cryptates.
12. (twice amended) A method as claimed in claim 24, wherein said chelate complex is a complex of a chelant selected from compounds of formulae (I), (II), (III), (IV), (V) and (VI):

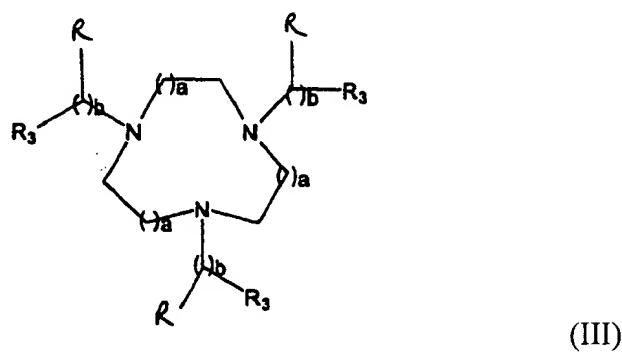


where each a independently represents an integer between 1 and 3, each R independently represents hydrogen or hydroxy and each R₁ independently represents a carboxylate, phosphate, thioacid, thiol, amino alkoxide or hydroxy group;

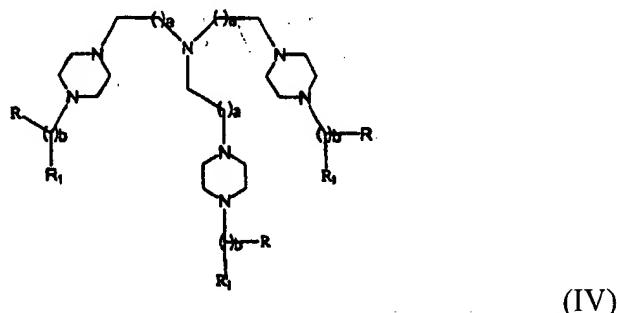


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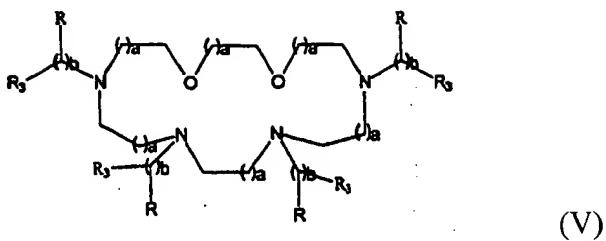
where a and R₁ are as hereinbefore defined and each R₂ independently represents hydrogen, C₁₋₆ alkyl or aryl, with the proviso that R₂ is absent when the double bond is present on the same nitrogen;



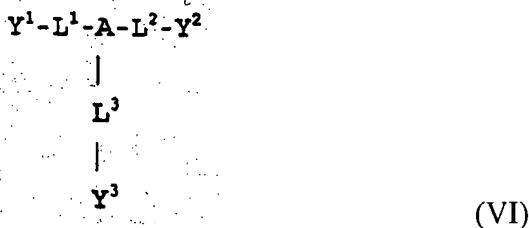
where a, R and R₂ are as hereinbefore defined, b is an integer between 0-3 and each R₃ independently represents R₁, NR-NR₂-COO[⊖], or N=N-COO[⊖] when b is positive or each R₃ independently represents N=CH-COO[⊖] or NR₂-CH₂-COO[⊖];



where a, b, R and R₁ are as hereinbefore defined;



where a, b, R and R₃ are as hereinbefore defined;



where A is N, CR₄, P, P=O, *cis,cis,cis*-1,3,5-trisubstituted-cyclohexane or an N,N',N"-triosubstituted-triaza 9 to 14 membered macrocyclic ring; L¹,L²,L³ are linker groups which are independently chosen from C₁₋₄ alkylene, C₄₋₈ cycloalkylene or C₄₋₈ o-arylene; Y¹,Y²,Y³ are independently chosen from -NH₂, -B(=O)OZ, -N=CR₅-B(=O)OZ, -NR₅-CR₆-(=O)OZ, -N[CR₆-B(=O)Q]₂ and -O-CR₆-B(=O)OZ where B is C or PR₆, each Q is independently -OZ or -NR₆, and Z is H or a counter-ion; each R₄ and R₅ group is independently chosen from H, C₁₋₅ alkyl, C₁₋₅ alkoxyalkyl, C₁₋₅ hydroxyalkyl, C₁₋₅ aminoalkyl, C₅₋₁₀ aryl or C₁₋₆ fluoroalkyl; R₆ is OH, C₁₋₆ alkyl, C₁₋₆ alkoxyalkyl, C₁₋₆ fluoroalkyl, C₁₋₁₀ alkoxy or C₅₋₁₀ aryl; with the proviso that at least one of Y¹, Y² and Y³ is -N=CR₅-B(=O)OZ.

13. (twice amended) The method as claimed in claim 23, wherein said contrast agent is conjugated to a biological vector capable of targeting said contrast agent to a desired region of the body.

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14. (once amended) The method as claimed in claim 13, wherein said biological vector is selected from the group consisting of an antibody, an antibody fragment, and an oligonucleotide binding motif.

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23. (new) A method of detecting regions with decreased vascular perfusion in a human or non-human animal subject which comprises

- a) administering to said subject an effective amount of a magnetic resonance imaging contrast agent comprising a physiologically tolerable Europium (II) compound having a first oxidation state and wherein said Europium (II) compound is oxidized *in vivo* to a Europium (III) compound having a second oxidation state and said oxidation states differ in relaxivity by a factor of at least 5, whereby contrast difference is enhanced in regions with decreased vascular perfusion in which conversion between said oxidation states occurs; and
- b) generating an image of said subject.

24. (new) The method as claimed in claim 23, wherein said Europium (II) compound is a chelate complex of Europium (II) or a physiologically tolerable salt thereof.

25. (new) The method as claimed in claim 23, wherein said oxidation states differ in relaxivity by a factor of at least 10.

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- 26. (new) The method as claimed in claim 23, wherein said oxidation states differ in relaxivity by a factor of at least 20.
- 27. (new) The method as claimed in claim 23, wherein said oxidation states differ in relaxivity by a factor of at least 100.
- 28. (new) The method as claimed in claim 23, wherein said contrast agent is conjugated to a macromolecule selected from the group consisting of proteins, polymers and liposomes.
- 29. (new) The method as claimed in claim 23, wherein said regions are tumours.
- 30. (new) The method as claimed in claim 23, wherein said regions are cardiac tissue.
- 31. (new) The method as claimed in claim 23, wherein said regions are in the brain.
- 32. (new) The method as claimed in claim 25, wherein the method is used in the evaluation of stroke.